(I)

AMENDMENTS TO THE CLAIMS:

Listing of Claims:

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This listing of claims will replace all prior versions of the claims and listing of the claims in the application:

1. (Currently Amended) A substituted tetracycline compound, wherein said compound is of the formula:

wherein:

X is CHC(R13Y'Y), CR6'R6, S. NR6, or O:

R² is hydrogen, alkyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, alkylsumino, arylalkyl, aryl, heterocyclic, heteroaromatic or a prodrug moiety;

R⁴ and R⁴ are each hydrogen, alkyl, alkenyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, alkylamino, arylalkyl, aryl, heterocyclic, heteroaromatic or a prodrug moiety;

R2', R3, R10, R11 and R12 are each hydrogen or a pro-drug moiety;

R5 is hydrogen, hydroxyl, or a prodrug moiety;

 R^6, R^6 , and R^8 are each independently hydrogen, alkyl, alkenyl, alkynyl, aryl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, alkylamino, arylalkyl, or halogen;

R7 is hydrogen, or NR7cC(=W')WR7a;

R8 is hydrogen;

 R^{13} is hydrogen, hydroxy, alkyl; alkenyl; alkynyl; alkynyl; alkylthio; alkylsulfinyl; alkylsulfonyl; alkylamino; or an arylalkyl;

Y' and Y are each independently hydrogen; halogen; hydroxyl; cyano, sulfhydryl; amino; alkyl; alkenyl; alkenyl; alkenyl; alkynyl; alkylsulfinyl; alkylsulfinyl; alkylsulfonyl; alkylamino; or an arylalkyl;

R9 is hydrogen, or NR9cC(=Z')ZR9a;

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Z is O:

Z' is O or S;

R²⁶ is unsubstituted C₂-C₁₀ alkyl, substituted alkyl, substituted or unsubstituted alkonyl, substituted or unsubstituted alkonyl, substituted or unsubstituted alkonyl, substituted or unsubstituted alkylsulfinyl, substituted or unsubstituted alkylsulfinyl, substituted or unsubstituted alkylsulfinyl, substituted or unsubstituted alkonylsulfinyl, substituted or unsubstituted alkonylsulfinyl, substituted or unsubstituted alkonylsulfinyl, substituted or unsubstituted alkylsulfino, substituted or unsubstituted arylalkyl, or substituted or unsubstituted heteroeyelic, substituted or unsubstituted or

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wherein said substituted alkyl is substituted with halogen, amino, hydroxyl, alkoxy, alkvayearbonyloxy, alkyloxycarbonyl, arylearbonyloxy, alkoxycarbonylamino, alkoxycarbonyloxy, aryloxycarbonyloxy, carboxylate, alkylearbonyl, alkylaminocarbonyl, arylalkyl aminocarbonyl, alkylaminocarbonyl, phosphate, aralkyl, phosphonato, phosphinato, cyano, acylamino, amido, imino, sulfhydryl, alkylthio, sulfate, arylthio, thiocarboxylate, alkylsulfinyl, sulfonato, sulfamoyl, sulfonamido, nitro, cyano, azido, heterocyclyl, alkylayl, aryl or heteroaryl:

further wherein said substituted alkenyl, substituted alkynyl, substituted alkoxy, substituted alkylsulfonyl, substituted alkylsulfonyl, substituted alkylsulfonyl, substituted arylsulfonyl, substituted arylsulfonamido, mitro, cyano, acido, heterocyclyl, alkylsulfinyl, sulfonato, sulfamoyl, sulfonamido, nitro, cyano, acido, heterocyclyl, alkylsulfinyl, sulfonato, sulfamoyl, sulfonato, sulfamoyl,

^{9e} is hydrogen, alkyl, alkenyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, arylsulfonyl, alkoxycarbonyl, arylcarbonyl, alkylamino, arylalkyl, aryl, heterocyclic or heteroaromatic:

W' is O or S:

R^{7a}, R^{7b}, R^{7c}, R^{7d}, and R^{7e} are each independently hydrogen, alkyl, alkenyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, arylsulfonyl, alkoxycarbonyl, arylcarbonyl, alkylamino, arylalkyl, aryl, heterocyclic, heteroaromatic, absent, or a prodrug moiety, and R^{7d} and R^{7c} may be linked to form a ring;

and pharmaceutically acceptable salts thereof, provided that R^{9} is not hydrogen when R^{7} is hydrogen.

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2. (Currently Amended) The compound of claim 1, wherein R^2 , R^2 , R^3 ,[[,,]] R^{10} , R^{11} , and R^{12} are each hydrogen.

- (Original) The compound of claim 2, wherein R⁴ and R⁴ are each alkyl.
- 4. (Currently Amended) The compound of claim 3, wherein \mathbb{R}^4 and $\mathbb{R}^{4'}$ are each methyl.
- (Canceled)
- 6. (Original) The compound of claim 4, wherein R⁵ is hydrogen.
- 7. (Original) The compound of claim 6, wherein X is CH₂, and R⁷ is hydrogen.
- 8. (Canceled)
- 9. (Previously Presented) The compound of claim 4, wherein ${\rm R}^5$ is hydroxyl and ${\rm X}$ is ${\rm CHR}^6.$
- 10. (Previously Presented) The compound of claim 9, wherein R⁶ is CH₃.
- 11. (Original) The compound of claim 1, wherein R⁹ is NR^{9c}C(=Z')ZR^{9a}.
- 12. (Original) The compound of claim 11, wherein R^{9c} is hydrogen.
- 13. (Original) The compound of claim 11, wherein Z' is oxygen.
- 14. (Original) The compound of claim 11, wherein Z' is sulfur.
- 15-18. (Canceled)
- (Currently Amended) The compound of claim 11, wherein R^{9a} is selected from the group consisting of substituted G₂-Ci₂-alkyl, <u>substituted or unsubstituted</u> alkynyl, <u>or</u> substituted or unsubstituted arvl-<u>arvalkely.</u> or-heteroarematic.
- 20. (Canceled)
- 21. (Currently Amended) The compound of claim 19, wherein said substituted G₂-G₄₉ alkyl is substituted with one or more substituents selected from the group consisting of alkoxycarbonyl, amino, arylcarbonyl, halogen, hydroxy, alkylamino, alkoxy, or aryl.
- 22. (Canceled)
- (Previously Presented) The compound of claim 19, wherein said substituted alkyl is substituted with an aryl group.
- 24. (Original) The compound of claim 23, wherein said aryl group is phenyl.
- 25. (Previously Presented) The compound of claim 19, wherein said substituted alkyl is substituted with one or more halogens.

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age -J-

26. (Currently Amended) The compound of elaim 24claim 25, wherein said halogen is bromine.

27-29. (Canceled)

- 30. (Currently Amended) The compound of elaim 19claim 1, wherein R^{9a} is substituted or unsubstituted aryl.
- (Original) The compound of claim 30, wherein said substituted or unsubstituted aryl is naphthyl.
- 32. (Currently Amended) The compound of elaim 30 claim 23, wherein said substituted or unsubstituted aryl is of the formulasaid substituted alkyl group is:

- 33. (Original) The compound of claim 30, wherein said substituted or unsubstituted aryl is phenyl.
- 34. (Currently Amended) The compound of claim 33, wherein said arylphenyl is substituted with one or more substituents selected from the group consisting of alkyl, alkenyl, alkynyl, aryl, alkoxy, aryloxy, alkylcarbonyl, arylcarbonyl, alkoxycarbonyl, aryloxycarbonyl, amido, halogen, nitro, azo, alkyl sulfonyl, and arylsulfonyl.
- 35. (Original) The compound of claim 34, wherein said substituent is alkyl.
- 36. (Original) The compound of claim 35, wherein said alkyl is unsubstituted.
- 37. (Original) The compound of claim 35, wherein said alkyl is methyl.
- 38. (Original) The compound of claim 35, wherein said alkyl is substituted with one or more halogens.
- 39. (Currently Amended) The compound of claim 34, wherein said substituent is methoxyalkoxy and further wherein said alkoxy is methoxy.
- (Original) The compound of claim 34, wherein said substituent is selected from the group consisting of alkylearbonyl, arylearbonyl, alkoxycarbonyl, aryloxycarbonyl, and amido.

41-55. (Canceled)

(Original) The compound of claim 1, wherein R⁷ is NR^{7c}C(=W')WR^{7a}.

- 57. (Original) The compound of claim 56, wherein R⁹ is hydrogen.
- 58. (Original) The compound of claim 57 wherein R^{7c} is hydrogen.
- 59. (Currently Amended) The compound of claim 57, wherein W' is oxygenO.
- 60. (Currently Amended) The compound of claim 57, wherein W' is sulfurS.
- 61. (Original) The compound of claims 59 or 60, wherein W is NR^{7b}.
- 62. (Currently Amended) The compound of claims 59 or 60, wherein W is oxygenO.
- 63. (Previously Presented) The compound of claim 57, wherein \mathbb{R}^{7a} is selected from the group consisting of alkyl, alkenyl, alkynyl, aryl, arylalkyl, and heteroaromatic.
- 64. (Original) The compound of claim 63, wherein \mathbb{R}^{7a} is substituted or unsubstituted alkyl.
- 65. (Original) The compound of claim 64, wherein said alkyl is substituted with an aryl group.
- 66. (Currently Amended) The compound of claim 63, wherein said substituted or unsubstituted R^{7a} is aryl and further wherein said aryl is phenyl.
- 67. (Currently Amended) The compound of claim 66, wherein said arylphenyl is substituted with one or more substituents selected from the group consisting of alkyl, alkenyl, alkynyl, aryl, alkoxy, aryloxy, arlylcarbonyl, arylcarbonyl, arkozerbonyl, aryloxycarbonyl, amido, halogen, nitro, azo, alkyl sulfonyl, and arylsulfonyl.
- (Original) The compound of claim 67, wherein said substituent is alkyl, alkoxy, or nitro.

69-81. (Canceled)

82. (Currently Amended) A pharmaceutical composition comprising a therapeutically effective amount of a substituted tetracycline compound and a pharmaceutically acceptable carrier, wherein said substituted tetracycline is of the formula:

$$\bigcap_{R^2}^{R^2} \bigcap_{Q \in \mathcal{Q}_{10}}^{R^2} \bigcap_{Q \in \mathcal{Q}_{$$

(I)

wherein:

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X is CHC(R13Y'Y), CR6'R6, S, NR6, or O;

R² is hydrogen, alkyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, alkylamino, arylalkyl, aryl, heterocyclic, heteroaromatic or a prodrug mojety;

R⁴ and R⁴ are each hydrogen, alkyl, alkenyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, alkylamino, arylalkyl, aryl, heterocyclic, heteroaromatic or a prodrug moiety;

R2', R3, R10, R11 and R12 are each hydrogen or a pro-drug moiety;

R5 is hydrogen, hydroxyl, or a prodrug moiety;

R⁶, R⁶', and R⁸ are each independently hydrogen, alkyl, alkenyl, alkynyl, aryl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, alkylsulfinyl, arylalkyl, or halogen;

R7 is hydrogen or NR7cC(=W1)WR7a;

R8 is hydrogen;

R¹³ is hydrogen, hydroxy, alkyl; alkenyl; alkynyl; alkoxy; alkylthio; alkylsulfinyl; alkylsulfonyl; alkylamino; or an arylalkyl;

Y' and Y are each independently hydrogen; halogen; hydroxyl; cyano, sulthydryl; amino; alkyl; alkenyl; alkynyl; alkoxy; alkylthio; alkylsulfinyl; alkylsulfonyl; alkylamino; or an arylalkyl;

R9 is hydrogen, or NR9cC(=Z')ZR9a:

Z is O;

Z' is O or S;

R²⁶ is unsubstituted C₃-C₁₀ alkyl, substituted alkyl, substituted or unsubstituted alkonyl, substituted or unsubstituted alkonyl, substituted or unsubstituted alkonyl, substituted or unsubstituted alkylsulfinyl, substituted or unsubstituted alkylsulfinyl, substituted or unsubstituted alkylsulfinyl, substituted or unsubstituted arylcarbonyl, substituted or unsubstituted arylcarbonyl, substituted or unsubstituted arylcarbonyl, substituted or unsubstituted arylalkyl-or substituted or unsubstituted aryl-aubstituted or unsubstituted arylsubstituted or unsubstituted arylsubstituted or unsubstituted arylsubstituted or unsubstituted beteroxylsubstituted bet

wherein said substituted alkyl is substituted with halogen, amino, hydroxyl, alkoxy, alkylcarbonyloxy, alkylcarbonyloxy, alkylcarbonyloxy, alkylcarbonyloxy, alkylcarbonyloxy, arylcarbonyloxy, arylcarbonyloxy, arylcarbonyloxy, arylcarbonyl, alkylcarbonyl, alkylcarbonyl, arylcarbonyl, alkylcarbonyl, alkylcarbonyl, alkylcarbonyl, alkylcarbonyl, alkylcarbonyl, alkylcarbonyl, alkylthiocarbonyl, alkylthiocarbonyl, alkylthiocarbonyl, phosphate, aralkyl, phosphonato, phosphinato, cyano, acylamino, amido, imino, sulfhydryl, alkylthio, ulitet, arylthio, thiocarboxylate, alkylsulfinyl, sulfonato, sulfamoyl, sulfonamido, nitro, cyano, azido, heterocyclyl, alkylaryl, aryl or heteroaryl;

further wherein said substituted alkenyl, substituted alkynyl, substituted alkynyl, substituted alkynyl, substituted alkylsulfonyl, substituted alkylsulfonyl, substituted alkylsulfonyl, substituted alkylsulfonyl, substituted

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arylsulfonyl, substituted alkoxycarbonyl, substituted arylcarbonyl, substituted-alkylamino; substituted-arylalkyl, or substituted aryl-substituted heteroeyelic, or-substituted heteroeryelic, alkoxy, alkynyl, hydroxyl, alkoxy, alkoxy, alkoxy, alkynyl, hydroxyl, alkoxy, alkoxycarbonyloxy, alkoxycarbonyl, alkoxy, aryloxycarbonyl, arylalkylaminoacarbonyl, alkylaminoacarbonyl, alkylaminoacarbonyl, alkylaminoacarbonyl, alkylaminoacarbonyl, alkylthiocarbonyl, phosphate, aralkyl, phosphonato, phosphinato, cyano, acylamino, amido, imino, sulfhydryl, alkylthio, sulfate, arylthio, thiocarboxylate, alkylsulfinyl, sulfonato, sulfamoyl, sulfonamido, nitro, cyano, azido, heteroeyelyl, alkylaryl, aryl or heteroaryl;

^{9e} is hydrogen, alkyl, alkenyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, arylsulfonyl, alkoxyearbonyl, arylcarbonyl, alkylamino, arylalkyl, aryl, heterocyclic or heteroaromatic:

W is CR7dR7e, NR7b or O:

W' is O or S:

 R^{7a} , R^{7b} , R^{7c} , R^{7d} , and R^{7e} are each independently hydrogen, alkyl, alkenyl, alkonyl, alkylsuffinyl, arylsulfonyl, alkoxycarbonyl, arylcarbonyl, alkylamino, arylalkyl, aryl, heterocyclic, heteroaromatic, absent, or a prodrug moiety, and R^{7d} and R^{7e} may be linked to form a ring:

and pharmaceutically acceptable salts thereof, provided that \mathbb{R}^9 is not hydrogen when \mathbb{R}^7 is hydrogen.

83-102. (Canceled)

103. (Currently Amended) The compound of claim 1, wherein said compound is

or a pharmaceutically acceptable salt thereof.

104. (Currently Amended) The compound of claim 1, wherein said compound is

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or a pharmaceutically acceptable salt thereof.

105. (Currently Amended) The compound of claim 1, wherein said compound is

or a pharmaceutically acceptable salt thereof.

106. (Currently Amended) The compound of claim 1, wherein said compound is

or a pharmaceutically acceptable salt thereof.

107. (Currently Amended) The compound of claim 1, wherein said compound is

or a pharmaceutically acceptable salt thereof.

108. (Previously Presented) A substituted tetracycline compound, wherein said compound is selected from the group consisting of

$$\begin{array}{c|c} & \text{CII}_1 & \text{GII} & \text{NoMe}_{2} \\ & \text{T} & \text{OII} \\ & \text{OII} & \text{OII} \\$$

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or a pharmaceutically acceptable salt thereof.

- 109. (Currently Amended) The compound of claim 1, wherein said compound is doxycycline 9-carbamic acid 9H-fluoren-9-yl methyl ester or a pharmaceutically acceptable salt thereof.
- 110. (Currently Amended) The compound of claim 1, wherein said compound is FMOC 9-amino doxycvcline or a pharmaceutically acceptable salt thereof.
- 111. (Currently Amended) The compound of claim 1, wherein said compound is 9-(4'-fluorophenyl) doxycycline carbamate or a pharmaceutically acceptable salt thereof.
- 112. (Currently Amended) The compound of claim 1, wherein said compound is 9-(4'-methoxyphenyl) doxycycline carbamate or a pharmaceutically acceptable salt thereof.

113-116. (Canceled)

- 117. (Currently Amended) The compound of claim 1, wherein said compound is 9-(2'-bromoethyl) doxycycline carbamate or a pharmaceutically acceptable salt thereof.
- 118. (Currently Amended) The compound of claim 1, wherein said compound is 7-(4'-methylphenyl) sancycline carbamate or a pharmaceutically acceptable sait thereof.
- 119. (Currently Amended) The compound of claim 1, wherein said compound is doxycycline 7-carbamic acid 7H-fluoren-7-yl methyl ester or <u>a</u> pharmaceutically acceptable salt thereof.
- 120. (Currently Amended) The compound of claim 1, wherein said compound is 7-(naphthyn-1-y1-naphthyn-1-y1) doxycycline urea or a pharmaceutically acceptable salt thereof.
- 121. (Currently Amended) The compound of claim 1, wherein said compound is 7-(3-methyl-1-butyl) doxycycline urea or a pharmaceutically acceptable salt thereof.
- 122. (Currently Amended) The compound of claim 1, wherein said compound is 7-phenyl doxycycline urea or a pharmaceutically acceptable salt thereof.
- 123. (Currently Amended) The compound of claim 1, wherein said compound is 7-t-butyl doxycycline urea or a pharmaceutically acceptable salt thereof.
- 124. (Currently Amended) The compound of claim 1, wherein said compound is 7-Fmoc amino doxycycline or a pharmaceutically acceptable salt thereof.

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- 125. (Currently Amended) The compound of claim 1, wherein said compound is 7-(4'-chloro-2-trifluoromethylphenyl) doxycycline urea or a pharmaceutically acceptable salt thereof.
- 126. (Currently Amended) The compound of claim 1, wherein said compound is 7-(4'-fluorophenyl) doxycycline carbamate or a pharmaceutically acceptable salt thereof.
- 127. (Currently Amended) The compound of claim 1, wherein said compound is 7-(4'-methoxyphenyl) doxycycline carbamate or a pharmaceutically acceptable salt thereof.
- 128. (Currently Amended) The compound of claim 1, wherein said compound is 7-BOC amino doxycycline or a pharmaceutically acceptable salt thereof.
- 129. (Currently Amended) The compound of claim 1, wherein said compound is 7(naphthyn-1-y1naphthyn-1-y1) doxycycline thiourea 5-propanoic acid ester or a
 pharmaceutically acceptable salt thereof.
- 130. (Currently Amended) The compound of claim 1, wherein said compound is doxycycline 7-thiocarbamic acid 7*H*-fluoren-7-yl methyl ester or <u>a pharmaceutically</u> acceptable sait thereof.
- 131. (Currently Amended) The compound of claim 1, wherein said compound is 7-(naphthyn-1-yl-naphthyn-1-yl) doxycycline thiourea or a pharmaceutically acceptable salt thereof.
- 132. (Currently Amended) The compound of claim 1, wherein said compound is 7-(3-methyl-1-butyl) doxycycline thiourea or a pharmaceutically acceptable salt thereof.
- 133. (Currently Amended) The compound of claim 1, wherein said compound is 7-phenyl amino doxycycline thiourea or a pharmaceutically acceptable salt thereof.
- 134. (Currently Amended) The compound of claim 1, wherein said compound is 7-tbutyl amino doxycycline thiourea or a pharmaceutically acceptable salt thereof.
- 135. (Currently Amended) The compound of claim 1, wherein said compound is 7-(4²-chloro-2²-trifluoromethylphenyl) doxycycline thiourea or <u>a</u>pharmaceutically acceptable salt thereof.
- 136. (Currently Amended) The compound of claim 1, wherein said compound is 7-(4'-fluorophenyl) doxycycline thiocarbamate or a pharmaceutically acceptable salt thereof.
- 137. (Currently Amended) The compound of claim 1, wherein said compound is 7-(4'-methoxyphenyl) doxycycline thiocarbamate or a pharmaceutically acceptable sait thereof.
- 138. (Currently Amended) The compound of claim 1, wherein said compound is 7-(nephthyn-l-ylnaphthyn-l-yl) doxycycline urea 5-propanoic acid ester or a pharmaceutically acceptable sait thereof.
- 139. (Currently Amended) A tetracycline compound, wherein said compound is selected from the group consisting of:

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9-neopentyl minocycline carbamate;

9-BOC amino doxycycline;

9-(n-pentyl) minocycline carbamate; 9-BOC amino minocycline carbamate:

9-(n-pentyl) minocycline carbamate;

9-prop-2'-enyl minocycline carbamate;

9-ethyl minocycline carbamate:

9-n-butyl minocycline carbamate

9-n-but-3-enyl minocycline carbamate; and

9-i-butyl minocycline carbamate; or a pharmaceutically acceptable salt thereof.

140. (Currently Amended) A substituted tetracycline compound, wherein said compound is of the formula:

wherein:

X is CHC(R13Y'Y),CR6'R6,S, NR6, or O;

 R^2 is hydrogen, alkyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, alkylamino, arylalkyl, aryl, heterocyclic, heteroaromatic or a prodrug moiety;

R⁴ and R⁴ are each hydrogen, alkyl, alkenyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, alkylamino, arylalkyl, aryl, heterocyclic, heteroaromatic or a prodrug moiety;

R2', R3, R10, R11 and R12 are each hydrogen or a pro-drug moiety;

R5 is hydrogen, hydroxyl, or a prodrug moiety;

 R^6, R^6 , and R^8 are each independently hydrogen, alkyl, alkenyl, alkynyl, aryl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, alkylamino, arylalkyl, or halogen;

R7 is dialkylamino;

R8 is hydrogen;

 R^{13} is hydrogen, hydroxy, alkyl; alkenyl; alkynyl; alkoxy; alkylthio; alkylsulfinyl; alkylsulfonyl; alkylamino; or an arylalkyl;

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Y' and Y are each independently hydrogen; halogen; hydroxyl; cyano, sulfhydryl; amino; alkyl; alkenyl; alkynyl; alkynyl; alkylsulfinyl; alkylsulfonyl; alkylamino; or an arylalkyl.

 R^9 is $NR^{9c}C(=Z')ZR^{9a}$:

Z is O:

Z' is O or S:

R^{9a} is unsubstituted or substituted C₃-C₁₀ alkyl, substituted or unsubstituted C₄-C₁₀ alkenyl, substituted or unsubstituted alkynyl, substituted or unsubstituted alkynyl, substituted or unsubstituted alkysulfinyl, substituted or unsubstituted alkylsulfinyl, substituted or unsubstituted alkylsulfonyl, substituted or unsubstituted arylcarbonyl, substituted or unsubstituted alkylsulfonyl, substituted or unsubstituted arylcarbonyl, substituted or unsubstituted arylcarbonyl, substituted or unsubstituted or unsubstituted or unsubstituted heteroarematic.

wherein said substituted $C_3 - C_{10}$ alkyl is substituted with halogen, hydroxyl, alkoxy, alkyloxycarbonyloxy, alkyloxycarbonyloxy, alkyloxycarbonyloxy, alkyloxycarbonyloxy, aryloxycarbonyloxy, aryloxycarbonyloxy, carboxyloxy, carboxyloxy, carboxyloxy, aryloxycarbonyl, alkylcarbonyl, alkylthiocarbonyl, phosphate, aralkyl, phosphonato, phosphinato, cyano, acylamino, amido, imino, sulfhydryl, alkylthio, sulfate, arylthio, thiocarboxylate, alkylsulfinyl, sulfonato, sulfamoyl, sulfonamido, nitro, cyano, acyldo, heteroevelyl, alkylaryl aryl or heteroaryl:

further wherein said substituted C_3 — C_{10} , alkenyl, substituted alkynyl, substituted alkynyl, substituted alkynyl, substituted alkyleulfinyl, substituted alkyleulfinyl, substituted alkyleulfinyl, substituted aryleulfonyl, substituted aryleulfonyloxy, substituted aryleulfonyloxy, alkyoxyentonyl, arylearbonyloxy, alkoxyentonyl, arylearbonyloxy, alkoxyentonyl, arylearbonyloxy, alkoxyentonyl, arylearbonyl, alkyleulfonyl, arylonyl, arylonyl

R^{5c} is hydrogen, alkyl, alkenyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, arylsulfonyl, alkyoxycarbonyl, arylearbonyl, alkylamino, arylalkyl, aryl, heterocyclic or heteroaromatic; and pharmaceutically acceptable salts thereof.

- 141. (Previously Presented) The compound of claim 140, wherein R^2 , R^2 , R^3 , R^{10} , R^{11} , and R^{12} are each hydrogen.
- 142. (Currently Amended) The compound of claim 140, wherein R^4 and $R^{4^{\prime}}$ are each methyl.
- 143. (Previously Presented) The compound of claim 140, wherein R⁵ is hydrogen.

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144. (Canceled)

- 145. (Currently Amended) The compound of elaim 144claim 140, wherein R^{9c} is hydrogen.
- 146. (Currently Amended) The compound of elaim 144claim 140, wherein Z' is oxygenQ.
- 147. (Currently Amended) The compound of claim 144claim 140, wherein Z' is sulfurS.
- 148. (Canceled)
- 149. (Currently Amended) The compound of elaim-144claim 140, wherein R^{9a} is substituted C₅-C₁₀ alkyl, <u>substituted or unsubstituted</u> alkynyl, <u>or substituted or unsubstituted</u> aryl-arylalkyl-or heteroaromatie.
- 150. (Currently Amended) The compound of claim 149, wherein said substituted C₅-C₁₀ alkyl is substituted with one or more substitutents selected from the group consisting of alkoxycarbonyl, arylcarbonyl, halogen, hydroxyl, alkoxy, or aryl.
- 151. (Currently Amended) The compound of elaim 149 claim 140, wherein R^{9a} is substituted or unsubstituted aryl.
- 152. (Previously Presented) The compound of claim 151, wherein said substituted or unsubstituted aryl is phenyl.
- 153. (Currently Amended) The compound of elaim 154claim 152, wherein said anylphenyl is substituted with one or more substituents selected from the group consisting of alkyl, alkenyl, alkynyl, aryl, alkoxy, aryloxy, alkylcarbonyl, arylatoponyl, alkoxycarbonyl, aryloxycarbonyl, amido, halogen, nitro, azo, alkyl sulfonyl, and arylsulfonyl.
- 154. (Currently Amended) A pharmaceutical composition comprising a therapeutically effective amount of a substituted tetracycline compound and a pharmaceutically acceptable carrier, wherein said substituted tetracycline is of the formula:

(I)

wherein:

X is CHC(R13Y'Y), CR6'R6, S, NR6, or O;

R² is hydrogen, alkyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, alkylamino, arylalkyl, aryl, heterocyclic, heteroaromatic or a prodrug moiety;

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R⁴ and R⁴ are each hydrogen, alkyl, alkenyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, alkylamino, arylalkyl, aryl, heterocyclic, heteroaromatic or a prodrug moiety;

R2', R3, R10, R11 and R12 and are each hydrogen or a pro-drug moiety;

R⁵ is hydrogen, hydroxyl, or a prodrug moiety;

 R^6, R^6 , and R^8 are each independently hydrogen, alkyl, alkenyl, alkynyl, aryl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, alkylamino, arylalkyl, or halogen;

R7 is dialkylamino;

R8 is hydrogen:

R¹³ is hydrogen, hydroxy, alkyl; alkenyl; alkynyl; alkoxy; alkylthio; alkylsulfinyl; alkylsulfonyl; alkylamino; or an arylalkyl;

Y' and Y are each independently hydrogen; halogen; hydroxyl; cyano, sulfhydryl; amino; alkyl; alkenyl; alkynyl; alkoxy; alkylthio; alkylsulfinyl; alkylsulfonyl; alkylamino; or an arylalkyl;

 R^9 is $NR^{9c}C(=Z')ZR^{9a}$;

Z is O;

Z' is O or S:

 R^{9a} is unsubstituted or substituted C_3 - C_{10} alkyl, substituted or unsubstituted C_4 - C_{10} alkenyl, substituted or unsubstituted alkynyl, substituted or unsubstituted alkynyl, substituted or unsubstituted alkysulfinyl, substituted or unsubstituted alkysulfinyl, substituted or unsubstituted alkysulfonyl, substituted or unsubstituted arylcarbonyl, substituted heteroarylcarbonyl, substituted or unsubstituted arylcarbonyl, substituted or unsubstituted arylcarbonyl, substituted or unsubstituted arylcarbon

wherein said substituted $\underline{C_3} \cdot \underline{C_{10}}$ alkyl is substituted with halogen, hydroxyl, alkoxy, alkylcarbonyloxy, alkylcarbonyl, arylcarbonyloxy, alkoxycarbonylamino, alkylcarbonyl, arylcarbonyloxy, aryloxycarbonyloxy, carboxylate, alkylcarbonyl, alkylaminocarbonyl, alkylaminocarbonyl, alkylcarbonyl, arylcarbonyl, arylcarbonyl, alkenylcarbonyl, alkoxycarbonyl, silyl, aminocarbonyl, alkylthiocarbonyl, phosphate, aralkyl, phosphonato, phosphinato, cyano, acylamino, amido, imino, sulfhydryl, alkylthio, ulifate, arylthio, thiocarboxylate, alkylsulfinyl, sulfonato, sulfamoyl, sulfonamido, nitro, cyano, azido, heterocyclyl, alkylaryl, aryl or heteroaryl;

further wherein said substituted C_0 — C_{10} alkenyl, substituted alkynyl, substituted alkysyl, substituted alkylsyl, substituted alkylsylfonyl; substituted arylsulfonyl, substituted heterocyclic, or substituted arylsulfonyl, by substituted arylsulfonyl, subs

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arylalkyl aminocarbonyl, alkenylaminocarbonyl, alkylcarbonyl, arylcarbonyl, aminoalkyl, arylalkylcarbonyl, alkenylcarbonyl, alkoxycarbonyl, silyl, aminocarbonyl, alkylthiocarbonyl, phosphate, aralkyl, phosphonato, phosphinato, cyano, acylamino, amido, imino, sulfhydryl, alkylthio, sulfate, arylthio, thiocarboxylate, alkylsulfinyl, sulfonato, sulfamoyl, sulfonamido, nitro, cyano, azido, heterocyclyl, alkylaryl, aryl or heteroaryl; and

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R^{5c} is hydrogen, alkyl, alkenyl, alkynyl, alkoxy, alkylthio, alkylsulfinyl, alkylsulfonyl, arylsulfonyl, alkoxycarbonyl, arylcarbonyl, alkylamino, arylalkyl, aryl, heterocyclic or heteroaromatic; and pharmaceutically acceptable salts thereof.

- 155. (New) The compound of claim 140, wherein said compound is minocycline 9-carbamic acid 9H-fluoren-9-yl methyl ester or a pharmaceutically acceptable salt thereof.
- 156. (New) The compound of claim 140, wherein said compound is FMOC 9-amino minocycline or a pharmaceutically acceptable salt thereof.
- 157. (New) The compound of claim 140, wherein said compound is 9-(4'- fluorophenyl) minocycline carbamate or a pharmaceutically acceptable salt thereof.
- 158. (New) The compound of claim 140, wherein said compound is 9-(4'-Methoxyphenyl) minocycline carbamate or a pharmaceutically acceptable salt thereof.